L Number	Hits	Search Text	DB	Time stamp
1	66643	peptidase or protease or proteinase	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/05/26 14:54
7	3178	(brain or neuronal) near2 ((app) or (amyloid precursor protein))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/05/26 14:55
13	11	(peptidase or protease or proteinase) near3 ((brain or neuronal) near2 ((app) or (amyloid precursor protein)))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/05/26 14:55

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NEWS 4	Aug 08	PHARMAMarketLetter(PHARMAML) - new on STN
NEWS 5	Aug 19	Aquatic Toxicity Information Retrieval (AQUIRE) now available on STN
NEWS 6	Aug 26	Sequence searching in REGISTRY enhanced
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NEWS 8	Sep 16	Experimental properties added to the REGISTRY file
NEWS 9	Sep 16	CA Section Thesaurus available in CAPLUS and CA
NEWS 10	Oct 01	CASREACT Enriched with Reactions from 1907 to 1985
NEWS 11	Oct 24	BEILSTEIN adds new search fields
NEWS 12	Oct 24	Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 13	Nov 18	DKILIT has been renamed APOLLIT
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NEWS 15	Dec 04	CSA files on STN
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NEWS 19	Jan 29	Simultaneous left and right truncation added to COMPENDEX, ENERGY, INSPEC
NEWS 20	Feb 13	CANCERLIT is no longer being updated
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NEWS 27	Mar 20	EVENTLINE will be removed from STN
NEWS 28	Mar 24	PATDPAFULL now available on STN
NEWS 29	Mar 24	Additional information for trade-named substances without structures available in REGISTRY
NEWS 30	Apr 11	Display formats in DGENE enhanced
NEWS 31	Apr 14	MEDLINE Reload
NEWS 32	Apr 17	Polymer searching in REGISTRY enhanced
NEWS 33	Apr 21	Indexing from 1947 to 1956 being added to records in CA/CAPLUS
NEWS 34	Apr 21	New current-awareness alert (SDI) frequency in WPIDS/WPINDEX/WPIX
NEWS 35	Apr 28	RDISCLOSURE now available on STN
NEWS 36	May 05	Pharmacokinetic information and systematic chemical names added to PHAR
NEWS 37	May 15	MEDLINE file segment of TOXCENTER reloaded
NEWS 38	May 15	Supporter information for ENCOMPPAT and ENCOMPLIT updated
NEWS 39	May 16	CHEMREACT will be removed from STN
NEWS 40	May 19	Simultaneous left and right truncation added to WSCA
NEWS 41	May 19	RAPRA enhanced with new search field, simultaneous left and right truncation

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  61 FILES SEARCHED...
L1     1143939 PEPTIDASE OR PROTEASE OR PROTEINASE

=> s (brain or neuronal) (2A) app
  29 FILES SEARCHED...
  59 FILES SEARCHED...
L2        2298 (BRAIN OR NEURONAL) (2A) APP

=> s l1 (3A) l2
  56 FILES SEARCHED...
L3          18 L1 (3A) L2

=> s (Brain or neuronal) (2a) (amyloid precursor protein)
  13 FILES SEARCHED...
  23 FILES SEARCHED...
  38 FILES SEARCHED...
  54 FILES SEARCHED...
  68 FILES SEARCHED...
L4        1200 (BRAIN OR NEURONAL) (2A) (AMYLOID PRECURSOR PROTEIN)

=> s l4 or l2
  56 FILES SEARCHED...
L5        3224 L4 OR L2

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PROCESSING COMPLETED FOR L6
L7 48 DUPLICATE REMOVE L6 (10 DUPLICATES REMOVED)

=> d l7 1-48 bib

L7 ANSWER 1 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 2002:937303 CAPLUS
DN 138:20443
TI Endocrine disruptor screening using DNA chips of endocrine disruptor-responsive genes
IN Kondo, Akihiro; Takeda, Takeshi; Mizutani, Shigetoshi; Tsujimoto, Yoshimasa; Takashima, Ryokichi; Enoki, Yuki; Kato, Ikunoshin
PA Takara Bio Inc., Japan
SO Jpn. Kokai Tokkyo Koho, 386 pp.
CODEN: JKXXAF
DT Patent
LA Japanese
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 2002355079	A2	20021210	JP 2002-69354	20020313
PRAI	JP 2001-73183	A	20010314		
	JP 2001-74993	A	20010315		
	JP 2001-102519	A	20010330		

L7 ANSWER 2 OF 48 USPATFULL
AN 2002:191539 USPATFULL
TI Full-length human cDNAs encoding potentially secreted proteins
IN Milne Edwards, Jean-Baptiste Dumas, Paris, FRANCE
Bougueleret, Lydie, Petit Lancy, SWITZERLAND
Jobert, Severin, Paris, FRANCE
PI US 2002102604 A1 20020801
AI US 2000-731872 A1 20001207 (9)
PRAI US 1999-169629P 19991208 (60)
US 2000-187470P 20000306 (60)
DT Utility
FS APPLICATION
LREP John Lucas, Ph.D., J.D., Genset Corporation, 10665 Srrento Valley Road, San Diego, CA, 92121-1609
CLMN Number of Claims: 29
ECL Exemplary Claim: 1
DRWN 5 Drawing Page(s)
LN.CNT 28061
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 3 OF 48 USPATFULL
AN 2002:32581 USPATFULL
TI Methods to treat alzheimer's disease
IN Hom, Roy, San Francisco, CA, UNITED STATES
Mamo, Shumeye S., Oakland, CA, UNITED STATES
Tung, Jay, Belmont, CA, UNITED STATES
Gailunas, Andrea, San Francisco, CA, UNITED STATES
John, Varghese, San Francisco, CA, UNITED STATES
Fang, Lawrence Y., Foster City, CA, UNITED STATES
PI US 2002019403 A1 20020214
AI US 2001-816876 A1 20010323 (9)
PRAI US 2000-191528P 20000323 (60)

DT Utility
FS APPLICATION
LREP MERCHANT & GOULD PC, P.O. BOX 2903, MINNEAPOLIS, MN, 55402-0903
CLMN Number of Claims: 63
ECL Exemplary Claim: 1
DRWN No Drawings
LN.CNT 8655
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 4 OF 48 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE 1
AN 2001:204410 BIOSIS
DN PREV200100204410
TI Expression of human brain carboxypeptidase B, a possible cleaving enzyme for beta-amyloid precursor protein, in peripheral fluids.
AU Matsumoto, Akira (1); Motozaki, Kenjiro; Seki, Tsuneyoshi; Sasaki, Ryohei; Kawabe, Tetsuya
CS (1) Department of Radiation Biophysics and Genetics, Kobe University School of Medicine, Kusunoki-cho chuo-ki 7-5-1, Kobe, 650-0017: amat@med.kobe-u.ac.jp Japan
SO Neuroscience Research, (March, 2001) Vol. 39, No. 3, pp. 313-317. print. ISSN: 0168-0102.
DT Article
LA English
SL English

L7 ANSWER 5 OF 48 WPINDEX (C) 2003 THOMSON DERWENT
AN 2000-687534 [67] WPINDEX
DNN N2000-508290 DNC C2000-209319
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders.
DC B04 D16 S03
IN MATSUMOTO, A
PA (MATS-I) MATSUMOTO A
CYC 92
PI WO 2000066717 A1 20001109 (200067)* JA 84p
RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW
W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
AU 2000043166 A 20001117 (200111)
EP 1179588 A1 20020213 (200219) EN
R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI
JP 2000615742 X 20021203 (200309)
ADT WO 2000066717 A1 WO 2000-JP2878 20000501; AU 2000043166 A AU 2000-43166 20000501; EP 1179588 A1 EP 2000-922936 20000501, WO 2000-JP2878 20000501; JP 2000615742 X JP 2000-615742 20000501, WO 2000-JP2878 20000501
FDT AU 2000043166 A Based on WO 200066717; EP 1179588 A1 Based on WO 200066717; JP 2000615742 X Based on WO 200066717
PRAI JP 1999-125169 19990430

L7 ANSWER 6 OF 48 USPATFULL
AN 1999:155781 USPATFULL
TI Arylsulfonamides as phospholipase A.sub.2 inhibitors
IN John, Varghese, San Francisco, CA, United States
Rydel, Russell E., Belmont, CA, United States
Thorsett, Eugene D., Moss Beach, CA, United States
PA Elan Pharmaceuticals, Inc., South San Francisco, CA, United States (U.S. corporation)
PI US 5994398 19991130

AI US 1996-766554 19961211 (8)
DT Utility
FS Granted
EXNAM Primary Examiner: Ford, John M.; Assistant Examiner: Kifle, Bruck
LREP Townsend & Townsend & Crew
CLMN Number of Claims: 40
ECL Exemplary Claim: 1
DRWN 5 Drawing Figure(s); 2 Drawing Page(s)
LN.CNT 1939
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 7 OF 48 CBNB COPYRIGHT 2003 EI
AN 15(45):64780 CBNB
TI Decade-long search yields elusive Alzheimer's enzyme.
SO Biotechnology Newswatch (1 Nov 1999), 1,3, (200-899 words)
ISSN: 0275-3685
DT Journal
LA English
PY 1999

L7 ANSWER 8 OF 48 USPATFULL
AN 1998:4424 USPATFULL
TI Identification of phospholipase A2 inhibitors in A.beta. peptide-mediated neurodegenerative disease
IN Rydel, Russell E., Belmont, CA, United States
Dappen, Michael S., San Bruno, CA, United States
PA Athena Neurosciences, Inc., San Francisco, CA, United States (U.S. corporation)
PI US 5707821 19980113
AI US 1995-476464 19950607 (8)
DT Utility
FS Granted
EXNAM Primary Examiner: Wax, Robert A.; Assistant Examiner: Lau, Kawai
LREP Duvall, Esq., Jean, Storella, Esq., JohnTownsend and Townsend and Crew
CLMN Number of Claims: 7
ECL Exemplary Claim: 1
DRWN 12 Drawing Figure(s); 9 Drawing Page(s)
LN.CNT 1580
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 9 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1998:83026 CAPLUS
DN 128:280072
TI A human brain proteolytic activity capable of cleaving natural .beta.-amyloid precursor protein is affected by its substrate glycoconjugates
AU Matsumoto, Akira; Matsumoto, Reiko; Enomoto, Taira; Itoh, Kyoko
CS Chuo-ku, Kusunoki-cho 7-5-1, Department of Radiation Biophysics and Genetics, Kobe University School of Medicine, Kobe, Japan
SO Neuroscience Letters (1998), 242(2), 109-113
CODEN: NELED5; ISSN: 0304-3940
PB Elsevier Science Ireland Ltd.
DT Journal
LA English

L7 ANSWER 10 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1997:246789 CAPLUS
DN 126:315931
TI Amyloid precursor proteins protect neurons of transgenic mice against acute and chronic excitotoxic injuries in vivo
AU Masliah, E.; Westland, C. E.; Rockenstein, E. M.; Abraham, C. R.; Mallory, M.; Veinberg, I.; Sheldon, E.; Mucke, L.
CS Departments of Neurosciences and Pathology, University of California at San Diego, La Jolla, CA, 92093-0624, USA

SO Neuroscience (Oxford) (1997), 78(1), 135-146
CODEN: NRSCDN; ISSN: 0306-4522
PB Elsevier
DT Journal
LA English

L7 ANSWER 11 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1996:304682 CAPLUS
DN 125:7373
TI A novel brain cysteine protease forms an SDS stable complex with the .beta.-amyloid precursor protein
AU Chang, Tien; Abraham, Carmela R.
CS School of Medicine, Boston University, Boston, MA, 02118, USA
SO Annals of the New York Academy of Sciences (1996), 777(Neurobiology of Alzheimers Disease), 183-188
CODEN: ANYAA9; ISSN: 0077-8923
PB New York Academy of Sciences
DT Journal
LA English

L7 ANSWER 12 OF 48 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.
AN 1996:452988 BIOSIS
DN PREV199699175344
TI Hydrolysis of amyloid precursor protein (**APP**) by a **brain** acid **protease**.
AU Malik, M. N.; Wen, G. Y.; Wisniewski, H. M.
CS New York State Inst. Basic Res. Dev. Disabilities, 1050 Forest Hill Road, Staten Island, NY 10314 USA
SO Neurobiology of Aging, (1996) Vol. 17, No. 4 SUPPL., pp. S104-S105.
Meeting Info.: Fifth International Conference on Alzheimer's Disease and Related Disorders Osaka, Japan July 24-29, 1996
ISSN: 0197-4580.
DT Conference
LA English

L7 ANSWER 13 OF 48 USPATFULL
AN 95:50068 USPATFULL
TI Detection of brain .alpha.1-antichymotrypsin
IN Johnson-Wood, Kelly, Belmont, CA, United States
Schenk, Dale, Pacifica, CA, United States
PA Athena Neurosciences, Inc., South San Francisco, CA, United States (U.S. corporation)
PI US 5422244 19950606
AI US 1992-880216 19920505 (7)
DT Utility
FS Granted
EXNAM Primary Examiner: Bidwell, Carol E.
CLMN Number of Claims: 26
ECL Exemplary Claim: 17
DRWN 3 Drawing Figure(s); 1 Drawing Page(s)
LN.CNT 1421
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 14 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1994:292870 CAPLUS
DN 120:292870
TI Generation of potentially amyloidogenic fragments from the beta-**amyloid precursor protein** by **brain** serine **proteases**
AU Martin, Brownwyn L.
CS Boston Univ., Boston, MA, USA
SO (1994) 342 pp. Avail.: Univ. Microfilms Int., Order No. DA9330150
From: Diss. Abstr. Int. B 1993, 54(6), 3048
DT Dissertation

LA English

L7 ANSWER 15 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1994:652968 CAPLUS
DN 121:252968
TI Increased neuronal .beta.-amyloid precursor protein expression in human temporal lobe epilepsy: association with interleukin-1.alpha. immunoreactivity
AU Sheng, Jin G.; Boop, Frederick A.; Mrak, Robert E.; Griffin, W. Sue T.
CS Arkansas Children's Hospital Reseach Center, Departments of Anatomy, Neurosurgery, Pathology and Pediatrics, Little Rock, AR, USA
SO Journal of Neurochemistry (1994), 63(5), 1872-9
CODEN: JONRA9; ISSN: 0022-3042
DT Journal
LA English

L7 ANSWER 16 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1995:47779 CAPLUS
DN 122:181691
TI Potential .beta.PP-processing proteinase activities from Alzheimer's and control brain tissues
AU Ladror, Uri S.; Wang, Gary T.; Klein, William L.; Holzman, Thomas F.; Krafft, Grant A.
CS Drug Design Delivery, Abbott Laboratories, Abbott Park, IL, 60064, USA
SO Journal of Protein Chemistry (1994), 13(4), 357-66
CODEN: JPCHD2; ISSN: 0277-8033
DT Journal
LA English

L7 ANSWER 17 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1995:390035 CAPLUS
DN 123:50670
TI Cleavage of fluorogenic substrates for APP-processing proteases by human brain extracts. Ca2+-substrate interaction is responsible for Ca2+ stimulation of the neural protease activity
AU Wang, Gary T.; Ladror, Uri S.; Holzman, Thomas F.; Klein, William L.; Krafft, Grant A.
CS Pharmaceutical Product Division, Abbott Laboratories, Abbott Park, IL, 60064, USA
SO Molecular and Chemical Neuropathology (1994), 23(2/3), 191-9
CODEN: MCHNEM; ISSN: 1044-7393
DT Journal
LA English

L7 ANSWER 18 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1994:241734 CAPLUS
DN 120:241734
TI .beta.-amyloid precursor protein fragments and lysosomal dense bodies are found in rat brain neurons after ventricular infusion of leupeptin
AU Hajimohammadreza, I.; Anderson, V. E. R.; Cavanagh, J. B.; Seville, M. P.; Nolan, C. C.; Anderton, B. H.; Leigh, P. N.
CS University Department of Neurology, London, SE5 8AF, UK
SO Brain Research (1994), 640(1-2), 25-32
CODEN: BRREAP; ISSN: 0006-8993
DT Journal
LA English

L7 ANSWER 19 OF 48 CAPLUS COPYRIGHT 2003 ACS DUPLICATE 2
AN 1993:601106 CAPLUS
DN 119:201106
TI Identification of a chymotrypsin-like mast cell protease in rat brain capable of generating the N-terminus of the Alzheimer amyloid .beta.-protein
AU Nelson, Robert B.; Siman, Robert; Iqbal, Mohamed A.; Potter, Huntington

CS Dep. Neurobiol., Harvard Med. Sch., Boston, MA, 02115, USA
SO Journal of Neurochemistry (1993), 61(2), 567-77
CODEN: JONRA9; ISSN: 0022-3042
DT Journal
LA English

L7 ANSWER 20 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1994:479550 CAPLUS
DN 121:79550
TI Studies on brain proteases capable of degrading the .beta. amyloid precursor protein
AU Abraham, Carmela R.; Papastoitsis, Gregory; Razzaboni, Bronwyn L.; Kanemaru, Kazutomi; Pietropaolo, Michael; Conn, Kelly-Jo; Meckelein, Barbara
CS Sch. Med., Boston Univ., Boston, MA, 02118, USA
SO Portland Press Proceedings (1993), 6(PROTEOLYSIS AND PROTEIN TURNOVER), 197-202
CODEN: POPPEF; ISSN: 0966-4068
DT Journal; General Review
LA English

L7 ANSWER 21 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1992:509327 CAPLUS
DN 117:109327
TI .beta.-Amyloid precursor protein cleavage by a membrane-bound protease
AU Sisodia, Sangram S.
CS Sch. Med., Johns Hopkins Univ., Baltimore, MD, 21205, USA
SO Proceedings of the National Academy of Sciences of the United States of America (1992), 89(13), 6075-9
CODEN: PNASA6; ISSN: 0027-8424
DT Journal
LA English

L7 ANSWER 22 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1993:405295 CAPLUS
DN 119:5295
TI Association and release of the amyloid protein precursor of Alzheimer's disease from chick brain extracellular matrix
AU Small, D. H.; Nurcombe, V.; Moir, R.; Michaelson, S.; Monard, D.; Beyreuther, K.; Masters, C. L.
CS Dep. Pathol., Univ. Melbourne, Parkville, 3052, Australia
SO Journal of Neuroscience (1992), 12(11), 4143-50
CODEN: JNRSDS; ISSN: 0270-6474
DT Journal
LA English

L7 ANSWER 23 OF 48 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.
AN 1992:296904 BIOSIS
DN BR43:9254
TI CO-DISTRIBUTION OF PROTEASE NEXIN-1 AND **PROTEASE** NEXIN-2-**APP**-SEC IN **BRAINS** OF NON-HUMAN PRIMATES.
AU WAGNER S L; VAN NOSTRAND W E; LAU A L; SUZUKI M; FARROW J S; BARTUS R; SCHUPPEK R; NGUYEN A; COTMAN C W; CUNNINGHAM D D
CS SALK INSTITUTE BIOTECHNOLOGY/INDUSTRIAL ASSOCIATES INC., LA JOLLA, CALIF. 92037.
SO KEYSTONE SYMPOSIUM ON ADVANCES IN UNDERSTANDING NEURODEGENERATIVE DISORDERS, BIG SKY, MONTANA, USA, MARCH 28-APRIL 4, 1992. J CELL BIOCHEM SUPPL. (1992) 0 (16 PART E), 213.
CODEN: JCBSD7.
DT Conference
FS BR; OLD
LA English

L7 ANSWER 24 OF 48 CAPLUS COPYRIGHT 2003 ACS

AN 1992:631521 CAPLUS
DN 117:231521
TI A calcium-stimulated serine protease from monkey brain degrades the .beta.-amyloid precursor protein
AU Razzaboni, Bronwyn L.; Papastoitsis, Gregory; Koo, Edward H.; Abraham, Carmela R.
CS Sch. Med., Boston Univ., Boston, MA, 02118, USA
SO Brain Research (1992), 589(2), 207-16
CODEN: BRREAP; ISSN: 0006-8993
DT Journal
LA English

L7 ANSWER 25 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1992:548472 CAPLUS
DN 117:148472
TI Two-way cleavage of .beta.-amyloid protein precursor by multicatalytic proteinase
AU Kojima, Shinichi; Omori, Motoko
CS Res. Inst., Sumitomo Pharm. Co., Osaka, 554, Japan
SO FEBS Letters (1992), 304(1), 57-60
CODEN: FEBLAL; ISSN: 0014-5793
DT Journal
LA English

L7 ANSWER 26 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1992:548494 CAPLUS
DN 117:148494
TI .alpha.2-Macroglobulin and other proteinase inhibitors do not interfere with the secretion of amyloid precursor protein in mouse neuroblastoma cells
AU De Strooper, Bart; Van Leuven, Fred; Van den Berghe, Herman
CS Cent. Hum. Genet., Univ. Leuven, Louvain, Belg.
SO FEBS Letters (1992), 308(1), 50-3
CODEN: FEBLAL; ISSN: 0014-5793
DT Journal
LA English

L7 ANSWER 27 OF 48 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.
AN 1992:245767 BIOSIS
DN BR42:116067
TI A SERINE PROTEASE FROM MONKEY AND ALZHEIMER'S BRAIN AND A CYSTEINE **PROTEASE** FROM ALZHEIMER'S **BRAIN** DEGRADE THE **AMYLOID PRECURSOR PROTEIN**.
AU ABRAHAM C R; RAZZABONI B; PAPASTOITSIS G
CS ARTHRITIS CENT., BOSTON UNIV. SCH. MED., BOSTON, MASS. 02118.
SO 21ST ANNUAL MEETING OF THE SOCIETY FOR NEUROSCIENCE, NEW ORLEANS, LOUISIANA, USA, NOVEMBER 10-15, 1991. SOC NEUROSCI ABSTR. (1991) 17 (1-2), 1105.
CODEN: ASNEE5.
DT Conference
FS BR; OLD
LA English

L7 ANSWER 28 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1991:119669 CAPLUS
DN 114:119669
TI A calcium-activated protease from Alzheimer's disease brain cleaves at the N-terminus of the amyloid .beta.-protein
AU Abraham, Carmela R.; Driscoll, James; Potter, Huntington; Van Nostrand, William E.; Tempst, Paul
CS Sch. Med., Boston Univ., Boston, MA, 02118, USA
SO Biochemical and Biophysical Research Communications (1991), 174(2), 790-6
CODEN: BBRCA9; ISSN: 0006-291X
DT Journal

LA English

L7 ANSWER 29 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1991:79412 CAPLUS
DN 114:79412
TI Proteolytic cleavage of the Alzheimer's disease amyloid A4 precursor protein
AU Ishiura, Shoichi
CS Natl. Inst. Neurosci., NCNP, Tokyo, 187, Japan
SO Journal of Neurochemistry (1991), 56(2), 363-9
CODEN: JONRA9; ISSN: 0022-3042
DT Journal; General Review
LA English

L7 ANSWER 30 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1993:122247 CAPLUS
DN 118:122247
TI An immunohistochemical study of amyloid precursor proteins containing the proteinase inhibitor region (APPI) in the human neocortex
AU Nakamura, Shinichi; Akiguchi, Ichiro; Kimura, Jun; Nakamura, Shigenobu; Tokushima, Yasuo; Kitaguchi, Nobuya; Takahashi, Yasuyuki; Shiojiri, Satoshi
CS Fac. Med., Kyoto Univ., Kyoto, Japan
SO International Congress Series (1991), 999(.beta.-Amyloid Precursor Proteins Neurotransm. Funct.), 26-37
CODEN: EXMDA4; ISSN: 0531-5131
DT Journal
LA English

L7 ANSWER 31 OF 48 BIOTECHABS COPYRIGHT 2003 THOMSON DERWENT AND ISI
AN 1990-12043 BIOTECHABS
TI Monoclonal antibody preparation;
against human brain amyloid precursor protein; mouse hybridoma construction; potential application in Alzheimer disease diagnosis
PA Asahi-Chem.
PI JP 02138995 28 May 1990
AI JP 1988-290626 17 Nov 1988
PRAI JP 1988-201998 15 Aug 1988; JP 1987-291404 18 Nov 1987
DT Patent
LA Japanese
OS WPI: 1990-206694 [27]

L7 ANSWER 32 OF 48 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.DUPLICATE 4
AN 1991:55121 BIOSIS
DN BA91:33402
TI ALZHEIMER AMYLOID BETA-PROTEIN PRECURSOR IN SPERM DEVELOPMENT.
AU SHOJI M; KAWARABAYASHI T; HARIGAYA Y; YAMAGUCHI H; HIRAI S; KAMIMURA T; SUGIYAMA T
CS DEP. NEUROL., GUNMA UNIV. SCH. MED., MAEBASHI, GUNMA 371, JPN.
SO AM J PATHOL, (1990) 137 (5), 1027-1032.
CODEN: AJPAA4. ISSN: 0002-9440.
FS BA; OLD
LA English

L7 ANSWER 33 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1990:609064 CAPLUS
DN 113:209064
TI Activity and expression of amyloid precursor proteins possessing proteinase inhibitor regions
AU Kitaguchi, Nobuya; Shiojiri, Satoshi; Takahashi, Yasuyuki; Tokushima, Yasuo
CS Bio-Sci. Lab., Asahi Chem. Ind. Co. Ltd., Fuji, 416, Japan
SO Shinkei Kenkyu no Shinpo (1990), 34(3), 409-21

CODEN: SKNSAF; ISSN: 0001-8724
DT Journal; General Review
LA Japanese

L7 ANSWER 34 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1991:604994 CAPLUS
DN 115:204994
TI Protease inhibitor and cholinergic system in Alzheimer's disease
AU Nakamura, Shigenobu; Tanaka, Seigo; Araki, Wataru; Tsuji, Teruyuki; Kawashima, Shingo; Shiojiri, Satoshi; Takahashi, Yasuyuki; Kitaguchi, Nobuya; Ito, Hirataka
CS Fac. Med., Kyoto Univ., Kyoto, Japan
SO Advances in Behavioral Biology (1990), 38A(Basic, Clin., Ther. Aspects Alzheimer's Parkinson's Dis., Vol. 1), 41-6
CODEN: ADBBBW; ISSN: 0099-6246
DT Journal
LA English

L7 ANSWER 35 OF 48 BIOTECHABS COPYRIGHT 2003 THOMSON DERWENT AND ISI
AN 1989-05725 BIOTECHABS
TI Human senile plaque amyloid precursor protein and DNA; protease-inhibitor gene cloning from cell culture and expression in Escherichia coli; DNA sequence; DNA probe construction
PA Asahi-Chem.
PI EP 304013 22 Feb 1989
AI EP 1988-113283 16 Aug 1988
PRAI JP 1988-125660 25 May 1988; JP 1987-203298 15 Aug 1987
DT Patent
LA English
OS WPI: 1989-055458 [08]

L7 ANSWER 36 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1990:53245 CAPLUS
DN 112:53245
TI Tissue-specific expression of three types of .beta.-protein precursor mRNA: enhancement of protease inhibitor-harboring types in Alzheimer's disease brain
AU Tanaka, Seigo; Shiojiri, Satoshi; Takahashi, Yasuyuki; Kitaguchi, Nobuya; Ito, Hirataka; Kameyama, Masakuni; Kimura, Jun; Nakamura, Shigenobu; Ueda, Kunihiro
CS Fac. Med., Kyoto Univ., Kyoto, 606, Japan
SO Biochemical and Biophysical Research Communications (1989), 165(3), 1406-14
CODEN: BBRCA9; ISSN: 0006-291X
DT Journal
LA English

L7 ANSWER 37 OF 48 JICST-EPlus COPYRIGHT 2003 JST
AN 890550938 JICST-EPlus
TI Molecular genetics of neurological disorders. Molecular biological research on Alzheimer's disease.
AU KITAGUCHI NOBUYA; TAKAHASHI YASUYUKI
CS Asahikaseikogyo Seikagakuken
SO Shinkei Kenkyu no Shinpo (Advances in Neurological Sciences), (1989) vol. 33, no. 4, pp. 571-582. Journal Code: Z0693A (Fig. 8, Tbl. 1, Ref. 52)
ISSN: 0001-8724
CY Japan
DT Journal; General Review
LA Japanese
STA New

L7 ANSWER 38 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1989:207058 CAPLUS
DN 110:207058

TI Identification of different .beta. amyloid cDNAs cloned from the brain of a patient with sporadic Alzheimer's disease
AU Octave, J. N.; De Sauvage, F.; Macq, A. F.; Maloteaux, J. M.; Laterre, E. C.
CS Lab. Neurochim., Univ. Cathol. Louvain, Brussels, B-1200, Belg.
SO Neurochemistry International (1989), 14(2), 163-6
CODEN: NEUIDS; ISSN: 0197-0186
DT Journal
LA English

L7 ANSWER 39 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAB11461 Protein DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501
PRAI JP 1999-125169 19990430
DT Patent
LA Japanese
OS 2000-687534 [67]
DESC Human brain carboxypeptidase B protein SEQ ID NO 9.

L7 ANSWER 40 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAB11460 Protein DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501
PRAI JP 1999-125169 19990430
DT Patent
LA Japanese
OS 2000-687534 [67]
DESC Human brain carboxypeptidase B protein SEQ ID NO 5.

L7 ANSWER 41 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAB11459 Protein DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501
PRAI JP 1999-125169 19990430
DT Patent
LA Japanese
OS 2000-687534 [67]
DESC Human brain carboxypeptidase B protein SEQ ID NO 4.

L7 ANSWER 42 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAB11458 Protein DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501

PRAI JP 1999-125169 19990430
DT Patent
LA Japanese
OS 2000-687534 [67]
DESC Human brain carboxypeptidase B protein SEQ ID NO 3.

L7 ANSWER 43 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAB11457 Protein DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501
PRAI JP 1999-125169 19990430
DT Patent
LA Japanese
OS 2000-687534 [67]
CR N-PSDB: AAC81962
DESC Human brain carboxypeptidase B protein.

L7 ANSWER 44 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAC81965 DNA DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501
PRAI JP 1999-125169 19990430
DT Patent
LA Japanese
OS 2000-687534 [67]
DESC Human brain carboxypeptidase B DNA primer SEQ ID NO 8.

L7 ANSWER 45 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAC81964 DNA DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501
PRAI JP 1999-125169 19990430
DT Patent
LA Japanese
OS 2000-687534 [67]
DESC Human brain carboxypeptidase B DNA primer SEQ ID NO 7.

L7 ANSWER 46 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAC81963 DNA DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501
PRAI JP 1999-125169 19990430
DT Patent
LA Japanese

OS 2000-687534 [67]
DESC Human brain carboxypeptidase B DNA primer SEQ ID NO 6.

L7 ANSWER 47 OF 48 DGENE (C) 2003 THOMSON DERWENT
AN AAC81962 cDNA DGENE
TI Human brain carboxypeptidase B isolated from the hippocampus useful for screening agents for the treatment of Alzheimer's and other brain disorders -
IN Matsumoto A
PA (MATS-I) MATSUMOTO A.
PI WO 2000066717 A1 20001109 84p
AI WO 2000-JP2878 20000501
PRAI JP 1999-125169 19990430
DT Patent
LA Japanese
OS 2000-687534 [67]
CR P-PSDB: AAB11457
DESC Human brain carboxypeptidase B cDNA.

L7 ANSWER 48 OF 48 INVESTEXT COPYRIGHT 2003 TFS

AN 91:419901 INVESTEXT(tm) REPORT NUMBER:1145046
PGNO PAGE 7 OF 14
DN 1145046
TI Cephalon, Inc. - Company Report
AU Siegel, J.G.
CS HAMBRECHT & QUIST INCORPORATED; NEW YORK
CSR MID-ATLANTIC/MIDDLE ATLANTIC STATES; UNITED STATES OF AMERICA; NORTH AMERICA
CSTY Financial center investment bank-broker
PD 9 Oct 1991
DT COMPANY REPORT
FS Text Page; COMPANY REPORT
WC 436

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=> d l7 7 9 11 12 14 16 17 19 20 21 25 27 2829 33 bib ab
48 ANSWERS ARE AVAILABLE. SPECIFIED ANSWER NUMBER EXCEEDS ANSWER SET SIZE
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bib ab
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Enter an answer number, Example: 10
several answer numbers, Example: 3,7,10
a range of answer numbers, Example: 5-10
or a combination of these. Example: 3,7,9-10,15
ENTER ANSWER NUMBER OR RANGE (1):7 9 11 12 14 16 17 19 20 21 25 27 28 29 33

L7 ANSWER 7 OF 48 CBNB COPYRIGHT 2003 EI
AN 15(45):84780 CBNB
TI Decade-long search yields elusive Alzheimer's enzyme.
SO Biotechnology Newswatch (1 Nov 1999), 1,3, (200-899 words)
ISSN: 0275-3685
DT Journal
LA English
PY 1999
AB Amgen researchers have discovered the gene responsible for producing betasecretase. This protease enzyme starts the development of Alzheimer's disease by cutting the **amyloid precursor protein** of healthy **brain** cells. A 2nd **protease** enzyme, gamma-secretase then cuts it again, causing a build up of beta-amyloid plaque in the brain. The Amgen team has not only identified the gene but also used it to produce beta-secretase and replicate the enzyme's amyloid precursor protein cutting action. It seems that a protease inhibitor (like those used against Aids) could be effective in treating Alzheimer's disease. Furthermore a single protease inhibitor might suffice. However it will be a long time before any new treatment reaches the market. There are still doubts about the development of Alzheimer's disease and the role gamma- secretase plays in it. The results of SmithKline Beecham's research into this enzyme should be presented at the annual Society for Neuroscience meeting. US biotechnology company Elan Pharmaceuticals has already patented a chemical which it says is beta-secretase, although this is not the same as Amgen's. Amgen itself has refused to comment on whether it is seeking to patent its discovery.

L7 ANSWER 9 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1998:83026 CAPLUS
DN 128:280072
TI A human brain proteolytic activity capable of cleaving natural .beta.-amyloid precursor protein is affected by its substrate glycoconjugates
AU Matsumoto, Akira; Matsumoto, Reiko; Enomoto, Taira; Itoh, Kyoko
CS Chuo-ku, Kusunoki-cho 7-5-1, Department of Radiation Biophysics and Genetics, Kobe University School of Medicine, Kobe, Japan
SO Neuroscience Letters (1998), 242(2), 109-113
CODEN: NELED5; ISSN: 0304-3940
PB Elsevier Science Ireland Ltd.
DT Journal
LA English
AB Human brain proteins were partially purified by using arginine-Sepharose 4B affinity chromatog., which traps proteins having an affinity to certain groups of arginine residue, such as serine proteases and zymogens. Bound proteins were analyzed for binding and cleavage related to the brain .beta.-amyloid precursor protein (APP). They were then further sepd. and isolated using a preparative gel system having a liq.-phase collection

app., using a non-denaturing gel system. Each fractionated protein was also analyzed for the above activity using natural APP. Among these, we found several fractions that bind preferentially to APP treated with chondroitinase ABC but not to intact APP, and that also generate particular .beta.-amyloid contg. C-terminal peptides of APP via proteolysis. Our results suggest that sulfated glycoconjugates attached to APP play a role in the substrate specificity of APP for proteases, and also that the nature of natural APP processing mechanisms in vivo is very complex.

L7 ANSWER 11 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1996:304682 CAPLUS
DN 125:7373
TI A novel brain cysteine protease forms an SDS stable complex with the .beta.-amyloid precursor protein
AU Chang, Tien; Abraham, Carmela R.
CS School of Medicine, Boston University, Boston, MA, 02118, USA
SO Annals of the New York Academy of Sciences (1996), 777(Neurobiology of Alzheimers Disease), 183-188
CODEN: ANYAA9; ISSN: 0077-8923
PB New York Academy of Sciences
DT Journal
LA English
AB Alzheimer's disease (AD) brain accumulates .beta.-protein (A.beta.) a peptide proteolytically derived from the .beta.-amyloid precursor protein (APP). The abnormal prodn. and aggregation of A.beta. have been implicated in the pathogenesis of the disease. The mechanism of prodn. of A.beta. in vivo is not yet clear; but endoproteases capable of degrading APP are likely to be involved in the process. The authors have isolated a protease from AD brain by following its activity in digesting a synthetic peptide of 10 amino acids derived from the APP sequence flanking the N-terminus of A.beta.. The protease was purified by a fractionation scheme including ammonium sulfate pptn. and column chromatog. using hydrophobic interaction, anion exchange, affinity, hydroxyapatite and size exclusion gels. The purity of the final product was assessed on a silver stained SDS gel by the presence of a single band. Microsequencing was performed following trypsin digestion of the sample. Internal peptide sequences were found to have sequence homol. to cysteine proteases in the database. The enzyme requires DTT for activity and can be inhibited by specific inhibitors of cysteine but not serine proteases. The purified enzyme has a pI of 5.0 and a native tetrameric structure with subunits of 48 kDa each. The enzyme is capable of digesting APP and generating a short peptide recognizable by antibodies specific to the C-terminus of APP. Interestingly, the purified protease also forms heat- and SDS-stable complexes with APP.

L7 ANSWER 12 OF 48 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.
AN 1996:452988 BIOSIS
DN PREV199699175344
TI Hydrolysis of amyloid precursor protein (**APP**) by a **brain acid protease**.
AU Malik, M. N.; Wen, G. Y.; Wisniewski, H. M.
CS New York State Inst. Basic Res. Dev. Disabilities, 1050 Forest Hill Road, Staten Island, NY 10314 USA
SO Neurobiology of Aging, (1996) Vol. 17, No. 4 SUPPL., pp. S104-S105.
Meeting Info.: Fifth International Conference on Alzheimer's Disease and Related Disorders Osaka, Japan July 24-29, 1996
ISSN: 0197-4580.
DT Conference
LA English

L7 ANSWER 14 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1994:292870 CAPLUS
DN 120:292870

TI Generation of potentially amyloidogenic fragments from the beta-**amyloid precursor protein** by **brain** serine **proteases**
AU Martin, Brownwyn L.
CS Boston Univ., Boston, MA, USA
SO (1994) 342 pp. Avail.: Univ. Microfilms Int., Order No. DA9330150 From: Diss. Abstr. Int. B 1993, 54(6), 3048
DT Dissertation
LA English
AB Unavailable

L7 ANSWER 16 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1995:47779 CAPLUS
DN 122:181691
TI Potential .beta.PP-processing proteinase activities from Alzheimer's and control brain tissues
AU Ladror, Uri S.; Wang, Gary T.; Klein, William L.; Holzman, Thomas F.; Krafft, Grant A.
CS Drug Design Delivery, Abbott Laboratories, Abbott Park, IL, 60064, USA
SO Journal of Protein Chemistry (1994), 13(4), 357-66
CODEN: JPCHD2; ISSN: 0277-8033
DT Journal
LA English
AB Fluorogenic peptide substrates designed to encompass the reported .alpha.-secretory and amyloidogenic cleavage sites of the amyloid-.beta. precursor protein (.beta.PP) were used to analyze proteinase activities in brain exts. from control patients and those with Alzheimer's disease (AD). Activity against the secretory substrate at pH 7.5 in control and AD brains produced a major endopeptidase cleavage at the Lys687-Leu688 bond (.beta.PP770 numbering), consistent with the .beta.PP secretase cleavage. Activity in control brains against the amyloidogenic substrate at pH 7.5 produced one cleavage at the Ala673-Glu674 bond, two residues C-terminal to the amyloidogenic Met-Asp site. However, in three of four AD brains, the major cleavage was at the Asp-Ala bond, one residue from the amyloidogenic site. Both endopeptidase and carboxypeptidase activities in AD brains were lower than in control brains. Proteinase activities against the secretory substrate had a major optimum at pH 3.0-4.0 and another at pH 6.0-7.5. Proteinase activities against the amyloidogenic substrate had a major optimum at or below pH 3.0 and another at pH 6.0. Using both substrates, activities at low pH were higher in AD brains than in controls, while at pH above 6.5, activities in control brains were higher than in AD. These results indicate that the levels of proteolytic enzymes in AD brains are altered relative to controls.

L7 ANSWER 17 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1995:390035 CAPLUS
DN 123:50670
TI Cleavage of fluorogenic substrates for APP-processing proteases by human brain extracts. Ca2+-substrate interaction is responsible for Ca2+ stimulation of the neural protease activity
AU Wang, Gary T.; Ladror, Uri S.; Holzman, Thomas F.; Klein, William L.; Krafft, Grant A.
CS Pharmaceutical Product Division, Abbott Laboratories, Abbott Park, IL, 60064, USA
SO Molecular and Chemical Neuropathology (1994), 23(2/3), 191-9
CODEN: MCHNEM; ISSN: 1044-7393
DT Journal
LA English
AB The proteases that cleave amyloid precursor protein (APP) leading to generation of amyloid A.beta. peptide are potential targets for therapeutical intervention of Alzheimer disease. The authors have been pursuing the identification and characterization of these proteases using as probes the fluorogenic substrates encompassing the cleavage sites of APP that were described recently (Wang, G. T.; Krafft, G. A., 1992). This

article describes results of expts. designed to examine the effect of Ca2+ on the cleavage of these substrates by human brain exts. Fluorogenic substrates encompassing either the N-terminal amyloidogenic cleavage site or the secretory cleavage site were synthesized in 5 formats with various peripheral residues. Incubation with exts. from normal brain tissue revealed that more neg. charged amyloidogenic substrates were less reactive and exhibited larger rate enhancement in the presence of Ca2+. The results imply that Ca2+ stimulation of substrate cleavage by brain proteases occurs primarily as a result of Ca2+-substrate interactions and caution against interpretations that invoke the involvement of Ca2+-stimulated proteases in A.beta. formation.

L7 ANSWER 19 OF 48 CAPLUS COPYRIGHT 2003 ACS DUPLICATE 2
AN 1993:601106 CAPLUS
DN 119:201106
TI Identification of a chymotrypsin-like mast cell protease in rat brain capable of generating the N-terminus of the Alzheimer amyloid .beta.-protein
AU Nelson, Robert B.; Siman, Robert; Iqbal, Mohamed A.; Potter, Huntington
CS Dep. Neurobiol., Harvard Med. Sch., Boston, MA, 02115, USA
SO Journal of Neurochemistry (1993), 61(2), 567-77
CODEN: JONRA9; ISSN: 0022-3042
DT Journal
LA English
AB Cleavage after Met596 of the .beta.-amyloid precursor protein to generate the N-terminus of the amyloid .beta.-protein indicates the activity of a proteinase having chymotrypsin-like specificity. A chymotrypsin-like proteinase is further implicated in Alzheimer's disease by the increased synthesis of the proteinase inhibitor .alpha.1-antichymotrypsin in pathol. affected brain regions and by the presence in the amyloid deposits of inactivated forms of .alpha.1-antichymotrypsin (indicating irreversible binding to a target chymotrypsin-like proteinase). In the present report, the authors purified from rat brain a chymotrypsin-like proteinase that (a) binds with high affinity to human .alpha.1-antichymotrypsin, (b) proteolytically generates a .beta.-protein-contg. C-terminal fragment from full-length recombinant human .beta.-amyloid precursor protein, and (c) selectively cleaves methoxysuccinyl-Glu-Val-Lys-Met-p-nitroanilide (a substrate modeling the proteinase recognition domain for the .beta.-protein N-terminal cleavage site). Amino acid sequences of tryptic fragments of the purified rat brain chymotrypsin-like proteinase indicate an identity with rat mast cell proteinase I. Moreover, the ontogeny and compartmentalization of rat brain chymotrypsin-like proteinase are consistent with those of connective tissue-type mast cells in the meningeal and intracortical perivasculature. Because these areas in human brain form extensive .beta.-amyloid deposits in Alzheimer's disease, Down's syndrome, and hereditary cerebral hemorrhage with amyloidosis of Dutch origin, the present findings suggest that a brain mast cell chymotrypsin-like proteinase may participate in generating perivascular .beta.-protein, which ultimately aggregates into .beta.-amyloid deposits.

L7 ANSWER 20 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1994:479550 CAPLUS
DN 121:79550
TI Studies on brain proteases capable of degrading the .beta. amyloid precursor protein
AU Abraham, Carmela R.; Papastoitsis, Gregory; Razzaboni, Bronwyn L.; Kanemaru, Kazutomi; Pietropaolo, Michael; Conn, Kelly-Jo; Meckelein, Barbara
CS Sch. Med., Boston Univ., Boston, MA, 02118, USA
SO Portland Press Proceedings (1993), 6(PROTEOLYSIS AND PROTEIN TURNOVER), 197-202
CODEN: POPPEF; ISSN: 0966-4068
DT Journal; General Review
LA English

AB A review and discussion with 35 refs. Several lines of evidence suggest that amyloid proteins may play a role in pathogenesis of Alzheimer's disease (AD). A no. of brain proteases have been isolated and characterized that have the potential to participate in the abnormal proteolytic processing of .beta. amyloid precursor protein seen in AD and, hence, form potentially important targets for therapeutic intervention.

L7 ANSWER 21 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1992:509327 CAPLUS
DN 117:109327
TI .beta.-Amyloid precursor protein cleavage by a membrane-bound protease
AU Sisodia, Sangram S.
CS Sch. Med., Johns Hopkins Univ., Baltimore, MD, 21205, USA
SO Proceedings of the National Academy of Sciences of the United States of America (1992), 89(13), 6075-9
CODEN: PNASA6; ISSN: 0027-8424
DT Journal
LA English
AB The principal component of brain amyloid plaques in Alzheimer disease is .beta.-amyloid protein, a .apprxeq.4-kDa peptide derived from amyloid precursor proteins. Amyloid precursor proteins are secreted after proteolytic cleavage of the .beta.-amyloid peptide. In cultured cells, amyloid precursor protein is cleaved on the plasma membrane by a membrane-bound endoprotease and the specificity of peptide bond hydrolysis is largely independent of the primary sequence of the precursor. The principal determinants of the cleavage appear to be an .alpha.-helical conformation and the distance (12-13 residues) of the hydrolyzed bond from the plasma membrane.

L7 ANSWER 25 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1992:548472 CAPLUS
DN 117:148472
TI Two-way cleavage of .beta.-amyloid protein precursor by multicatalytic proteinase
AU Kojima, Shinichi; Omori, Motoko
CS Res. Inst., Sumitomo Pharm. Co., Osaka, 554, Japan
SO FEBS Letters (1992), 304(1), 57-60
CODEN: FEBLAL; ISSN: 0014-5793
DT Journal
LA English
AB The .beta.-amyloid protein (.beta.-AP) derived from a .beta.-amyloid protein precursor (APP) is a hallmark of Alzheimer disease. The abundant generation of .beta.-AP suggests an abnormal processing of APP. The main APP-processing enzyme was purified from the rat brain and identified as a macroprotein-like multicatalytic proteinase. The purified enzyme cleaved the Gln15-Lys16 bond of .beta.-AP, but in the presence of Ca2+ the enzyme altered to cleave at the N-terminus of .beta.-AP to release the extracellular domain of .beta.-AP. Functional change in this multicatalytic proteinase may result in abnormal processing of APP in Alzheimer disease.

L7 ANSWER 27 OF 48 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.
AN 1992:245767 BIOSIS
DN BR42:116067
TI A SERINE PROTEASE FROM MONKEY AND ALZHEIMER'S BRAIN AND A CYSTEINE **PROTEASE** FROM ALZHEIMER'S **BRAIN** DEGRADE THE **AMYLOID PRECURSOR PROTEIN**.
AU ABRAHAM C R; RAZZABONI B; PAPASTOITSIS G
CS ARTHRITIS CENT., BOSTON UNIV. SCH. MED., BOSTON, MASS. 02118.
SO 21ST ANNUAL MEETING OF THE SOCIETY FOR NEUROSCIENCE, NEW ORLEANS, LOUISIANA, USA, NOVEMBER 10-15, 1991. SOC NEUROSCI ABSTR. (1991) 17 (1-2), 1105.
CODEN: ASNEE5.
DT Conference

FS BR; OLD
LA English

L7 ANSWER 28 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1991:119669 CAPLUS
DN 114:119669
TI A calcium-activated protease from Alzheimer's disease brain cleaves at the N-terminus of the amyloid .beta.-protein
AU Abraham, Carmela R.; Driscoll, James; Potter, Huntington; Van Nostrand, William E.; Tempst, Paul
CS Sch. Med., Boston Univ., Boston, MA, 02118, USA
SO Biochemical and Biophysical Research Communications (1991), 174(2), 790-6
CODEN: BBRCA9; ISSN: 0006-291X
DT Journal
LA English
AB Alzheimer's disease, Down's syndrome and to a lesser extent, normal aged brains exhibit abnormal extracellular deposits of amyloid. The major component of brain amyloid is the .beta.-protein, a 4Kd fragment of the larger B-protein precursor. The finding of the abnormally processed .beta.-protein and a protease inhibitor (.alpha.1-antichymotrypsin) in the amyloid deposits prompted the authors to search for proteases which may generate the .beta.-protein from its precursor. The presence and partial purifn. is reported of one such proteolytic activity from Alzheimer's brain. Normal physiol. C-terminal cleavage of the secreted form of the .beta.-protein precursor occurs in the middle of the .beta.-protein suggesting that the .beta.-protein accumulates due to an alternative degrdn. pathway. Apparently, the protease activity participates in this abnormal pathway.

L7 ANSWER 29 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1991:79412 CAPLUS
DN 114:79412
TI Proteolytic cleavage of the Alzheimer's disease amyloid A4 precursor protein
AU Ishiura, Shoichi
CS Natl. Inst. Neurosci., NCNP, Tokyo, 187, Japan
SO Journal of Neurochemistry (1991), 56(2), 363-9
CODEN: JONRA9; ISSN: 0022-3042
DT Journal; General Review
LA English
AB A review with 60 refs. Amyloid A4 protein (.beta.-protein) is deposited in the brain of patients with Alzheimer disease (AD) as one of the main components of the extracellular cerebrovascular amyloid and neurofibrillary tangles. It is derived from a precursor protein and its formation is a rate-limiting step for brain degeneration in AD together with proteolytic cleavage. The topog. distribution of the proteinase and its substrates in the AD brain is discussed.

L7 ANSWER 33 OF 48 CAPLUS COPYRIGHT 2003 ACS
AN 1990:609064 CAPLUS
DN 113:209064
TI Activity and expression of amyloid precursor proteins possessing proteinase inhibitor regions
AU Kitaguchi, Nobuya; Shiojiri, Satoshi; Takahashi, Yasuyuki; Tokushima, Yasuo
CS Bio-Sci. Lab., Asahi Chem. Ind. Co. Ltd., Fuji, 416, Japan
SO Shinkei Kenkyu no Shinpo (1990), 34(3), 409-21
CODEN: SKNSAF; ISSN: 0001-8724
DT Journal; General Review
LA Japanese
AB A review with 5 refs. Cerebral deposits of amyloid .beta. protein (.beta.-AP) comprising about 40 amino acids, as senile plaque core and vascular amyloid, are characteristic of Alzheimer's disease (AD). Sensitive assays by the polymerase chain reaction method have revealed

that small amts. of mRNA species coding for novel .beta.-AP precursor proteins (APP), APP714 and APP563 (secreted type of APP751), exist in human brain, as well as larger amts. of mRNAs for APP695, APP751 and APP770. APP751 and APP770 comprise Kunitz type trypsin inhibitor regions (APPI). Several reports have opened discussions on the physiol. role of APPs: their C-terminal fragments show neurotoxic activity, secreted forms of APP are trophic for proliferation of fibroblasts, and APPI comprised by APP751 stimulates cell division. APPI was found to inhibit trypsin and chymotrypsin most strongly and also to inhibit leukocyte elastase and several blood-coagulation proteinases, but not to inhibit urokinase or thrombin. Utilizing specific probes for each type of APP mRNAs in RNA blot anal., the authors have found an increase of APP751 (1.1.apprx.1.3 fold) and APP770 (about 2 fold) mRNA levels in several regions of AD brain compared with age-matched controls. A similar increase of the ratio of APP751 plus APP770 mRNA in AD brain is also reported and an increase of APP695 mRNA has been found in specific brain regions. There have been several reports on detection of APPI in cerebrospinal fluid (CSF) by immunoblot anal. Both sol. forms of APPs, with APPI (112 kDa) and without APPI (91 kDa), have been found to be more abundant in CSF of AD than in controls by using a monoclonal antibody against APP695. By using an antiserum against synthetic APPI peptide, 125 kDa and 58 kDa APP have been found in CSF. The proteinases and their inhibitors are important for the formation of .beta.-AP fragment from APP.

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